INTERNATIONAL SEARCH REPORT

International application No.

PCT/US04/19453

| A. CLASSIFICATION OF SUBJECT MATTER | | | | | | |
|---|--|--|--|--|--|--|
| IPC(7) : C07D 491/044; A61K 31/407 | | | | | | |
| US CL: 548/453; 514/421 According to International Patent Classification (IPC) or to both national classification and IPC | | | | | | |
| B. FIELDS SEARCHED | TOTAL GROSSITION OF DEC. | | | | | |
| | 1 (0 () | | | | | |
| Minimum documentation searched (classification system followed b U.S.: 548/453; 514/421 | y classification symbols) | | | | | |
| Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched | | | | | | |
| Electronic data base consulted during the international search (name of data base and, where practicable, search terms used) CAS ONLINE | | | | | | |
| C. DOCUMENTS CONSIDERED TO BE RELEVANT | | | | | | |
| Category * Citation of document, with indication, where a | ppropriate, of the relevant passages Relevant to claim No. | | | | | |
| A WO 02/47610 A2 (FENICAL et al) 20 June 2002 (2 | | | | | | |
| | | | | | | |
| Further documents are listed in the continuation of Box C. | See patent family annex. | | | | | |
| Special categories of cited documents: | "T" later document published after the international filing date or priority | | | | | |
| "A" document defining the general state of the art which is not considered to be of particular relevance | date and not in conflict with the application but cited to understand the principle or theory underlying the invention | | | | | |
| "E" earlier application or patent published on or after the international filing date | "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone | | | | | |
| "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) | "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination | | | | | |
| "O" document referring to an oral disclosure, use, exhibition or other means | being obvious to a person skilled in the art | | | | | |
| "P" document published prior to the international filing date but later than the "&" document member of the same patent family priority date claimed | | | | | | |
| Date of the actual completion of the international search Date of mailing of the international search report | | | | | | |
| 13 October 2004 (13.10.2004) 29 NOV 2004 | | | | | | |
| Name and mailing address of the ISA/US | Authorized officer | | | | | |
| Mail Stop PCT, Attn: ISA/US Commissioner for Patents | Fiona T. Powers DEBORAH A. THOMAS | | | | | |
| PARALEGAL SPE Alexandria, Virginia 22313-1450 Bacsimile No. (703) 305-3230 Telephone No. 703-308-1235 | | | | | | |

PATENT COOPERATION TREATY

From the INTERNATIONAL SEARCHING AUTHORITY

| y C | REC'D | 0 | 1 | DEC 2004 | |
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| PCT | | PCT | | | |

| Lisa A Haii | | nnich IID | | | | | |
|---|--|--|------------------------------|-----------------------|---------------------------------|------------|--|
| Gray, Gary, Ware & Freidenrich LLP 4365 Executive Drive, Suite 1100 | | WRITTEN OPINION OF THE | | | | | |
| San Diego, California 92121-2133 | | WRITTEN OPINION OF THE INTERNATIONAL SEARCHING AUTHORITY | | | | | |
| | | | | MILKIMII | J1(1111 D11111011111 0 1 1 1 | | |
| | | | | | (PCT Rule 43bis.1) | | |
| | | | | Date of mailing | nu o c | W 2004 | |
| | | | | (day/month/year) | | | |
| Applicant's | or agent's file re | eference | | FOR FURTHER | ACTION See paragraph 2 below | | |
| Internation | al application No. | | International filing date | (day/month/year) | Priority date (day/month/ye | ar) | |
| PCT/US04 | /19453 | | 18 June 2004 (18.06.20) | | 20 June 2003 (20.06.2003) | | |
| Internation | al Patent Classific | cation (IPC) | or both national classifica | tion and IPC | No. | | |
| IPC(7): C0 |)7D491/044; A61 | K 31/407 an | d US Cl.: 548/453; 514/4 | 21 | | | |
| Applicant | | | | • | | | |
| THE REG | ENTS OF THE U | JNIVERSIT | Y OF CALIFORNIA | | | | |
| 1. This o | pinion contains in | ndications re | lating to the following iter | ns: | | | |
| \boxtimes | Box No. I | Basis of the | e opinion | | | | |
| | Box No. II | Priority | | | | | |
| | Box No. III | Non-establ | ishment of opinion with re | egard to novelty, inv | entive step and industrial app | licability | |
| | Box No. IV | Lack of un | ity of invention | | | | |
| | Box No. V Reasoned statement under Rule 43bis.1(a)(i) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement | | | | industrial | | |
| | Box No. VI | Certain documents cited | | | | | |
| | Box No. VII Certain defects in the international application | | | | | | |
| | Box No. VIII | Certain ob | servations on the internati | onal application | | • | |
| 2. FUR | THER ACTIO | N | | | | | |
| If a demand for international preliminary examination is made, this opinion will be considered to be a written opinion of the International Preliminary Examining Authority ("IPEA") except that this does not apply where the applicant chooses an Authority other than this one to be the IPEA and the chosen IPEA has notified the International Bureau under Rule 66.1bis(b) that written opinions of this International Searching Authority will not be so considered. | | | | | | | |
| If this opinion is, as provided above, considered to be a written opinion of the IPEA, the applicant is invited to submit to the IPEA a written reply together, where appropriate, with amendments, before the expiration of 3 months from the date of mailing of Form PCT/ISA/220 or before the expiration of 22 months from the priority date, whichever expires later. | | | | | | | |
| For further options, see Form PCT/ISA/220. | | | | | | | |
| 3. For f | further details, se | e notes to Fo | orm PCT/ISA/220. | | | | |
| Name and mailing address of the ISA/ US | | | Authorized office | DEBORAH A | A. THOMAS | | |
| Mail Stop PCT, Attn: ISA/US Commissioner for Patents | | | Fiona T. Powe | rs PARALEGAL | SPECIALIST | | |
| P.O. Box 1450 | | | man a se | GROU | HOOD DENT | | |
| Alexandria, Virginia 22313-1450 Facsimile No. (703) 305-3230 | | | Telephone No. | /UD-3U0-1433 | • | | |

WRITTEN OPINION OF THE INTERNATIONAL SEARCHING AUTHORITY

International application No.

PCT/US04/19453

| Box No | . I Basis of this opinion |
|---------------------|---|
| | |
| 1. With r it was | regard to the language, this opinion has been established on the basis of the international application in the language in which filed, unless otherwise indicated under this item. |
| | This opinion has been established on the basis of a translation from the original language into the following language, which is the language of a translation furnished for the purposes of international search (under Rules 12.3 and 23.1(b)). |
| | regard to any nucleotide and/or amino acid sequence disclosed in the international application and necessary to the discountries, this opinion has been established on the basis of: |
| a. | type of material |
| | a sequence listing |
| | table(s) related to the sequence listing |
| b. | format of material |
| | in written format |
| | in computer readable form |
| c. | time of filing/furnishing |
| | contained in international application as filed. |
| | filed together with the international application in computer readable form. |
| | furnished subsequently to this Authority for the purposes of search. |
| | Infinited subsequently to this runnerty for the purposes of seaton. |
| 3. | In addition, in the case that more than one version or copy of a sequence listing and/or table relating thereto has been filed or furnished, the required statements that the information in the subsequent or additional copies is identical to that in the application as filed or does not go beyond the application as filed, as appropriate, were furnished. |
| 4. Additi | ional comments: |
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WRITTEN OPINION OF THE INTERNATIONAL SEARCHING AUTHORITY

Form PCT/ISA/237 (Box No. V) (January 2004)

T-4"

International application No. PCT/US04/19453

| Box No. V Reasoned statement under Rule 43 bis.1(a)(i) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement | | | | |
|---|-----------------------------------|---|--|--|
| 1. Statement | | | | |
| Novelty (N) | Claims | 1-14 | YES | |
| Troversy (Try | Claims | | NO | |
| | | | | |
| Inventive step (IS) | Claims | | YES | |
| | Claims | NONE | NO | |
| Industrial applicability (IA) | Claims | 114 | YES | |
| industrial applicability (1A) | | NONE | NO | |
| | | | | |
| 2. Citations and explanations: | | | | |
| Claims 1 to 14 meet the criteria set out in PCT Arti compounds having the structure (I), pharmaceutical disorder therewith. | icle 33(2)-(3), to compositions (| because the prior art does not teachereof or method of treating a r | ach or fairly suggest the claimed nammalian cell proliferative | |
| Claims 1 to 14 meet the criteria set out in PCT Artican be made or used in industry. | icle 33(4), and | thus have industrial applicability | because the subject matter claimed | |
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